

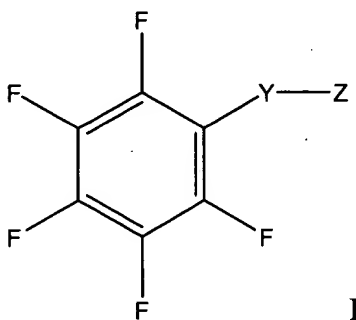
Amendments to the Claims:

This listing of claims will replace all prior versions, and listings of claims in the application:

Listing of Claims:

Please cancel claims 6, 7, 17, 20-26, 28, 29, 31, 35, 38, 42, 47, 50-53, 68, 74, 75, 77, 78, 81, 82, 85, 86, 88, 91, 93 and 94. Please amend claims 1 and 61 as indicated below.

a' 1. (amended) A pharmaceutical composition comprising a pharmaceutically acceptable excipient and a compound of formula 1:



or a pharmaceutically acceptable salt thereof, wherein:

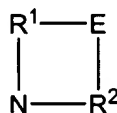
Y is -S(O)- or -S(O)₂-; and

Z is -NR¹R² or -OR³; wherein R¹ and R² are independently selected from hydrogen,

substituted or unsubstituted (C1-C10)alkyl,
substituted or unsubstituted (C1-C10)alkoxy,
substituted or unsubstituted (C3-C6)alkenyl,
substituted or unsubstituted (C2-C6)heteroalkyl,
substituted or unsubstituted (C3-C6)heteroalkenyl,
substituted or unsubstituted (C3-C6)alkynyl,
substituted or unsubstituted (C3-C8)cycloalkyl,
substituted or unsubstituted (C5-C7)cycloalkenyl,
substituted or unsubstituted (C5-C7)cycloalkadienyl,

a'
substituted or unsubstituted aryl,
substituted or unsubstituted aryloxy,
substituted or unsubstituted aryl-(C3-C8)cycloalkyl,
substituted or unsubstituted aryl-(C5-C7)cycloalkenyl,
substituted or unsubstituted aryloxy-(C3-C8)cycloalkyl,
substituted or unsubstituted aryl-(C1-C4)alkyl,
substituted or unsubstituted aryl-(C1-C4)alkoxy,
~~substituted or unsubstituted aryl-(C1-C4)heteroalkyl~~
substituted or unsubstituted aryl-(C3-C6)alkenyl,
substituted or unsubstituted aryloxy-(C1-C4)alkyl
substituted or unsubstituted aryloxy-(C2-C4)heteroalkyl,
substituted or unsubstituted heteroaryl,
substituted or unsubstituted heteroaryloxy,
substituted or unsubstituted heteroaryl-(C10-C4)alkyl,
substituted or unsubstituted heteroaryl-(C1-C4)alkoxy,
substituted or unsubstituted heteroaryl-(C1-C4)heteroalkyl,
substituted or unsubstituted heteroaryl-(C3-C6)alkenyl,
substituted or unsubstituted heteroaryloxy-(C1-C4)alkyl, and
substituted or unsubstituted heteroaryloxy-(C2-C4)heteroalkyl,

wherein R¹ and R² may be connected by a linking group E to give a substituent of the formula



wherein E represents a bond, (C1-C4) alkylene, or (C1-C4) heteroalkylene, and the ring formed by R¹, E, R² and the nitrogen atom contains no more than 8 atoms; and where R³ is a substituted or unsubstituted aryl or heteroaryl group, wherein said compound I has pharmacological activity.

2. (original) The composition of claim 1, wherein, in the compound of formula I, Y is SO₂ and

Z is NR^1R^2 ; wherein R^2 is optionally substituted aryl or optionally substituted heteroaryl.

a' 3. (original) The composition of claim 2, wherein R^1 is hydrogen or lower alkyl, R^2 is optionally substituted phenyl or optionally substituted pyridyl, and there is no linking group E between R^1 and R^2 .

4. (original) The composition of claim 3, wherein R^1 is hydrogen or methyl and R^2 is substituted phenyl, wherein the substituents on R^2 , ranging in number from one to four, are independently chosen from lower alkyl, hydroxy, lower alkoxy; amino optionally substituted with one or two lower alkyls, optionally substituted arylamino, optionally substituted heteroaryl amino, optionally substituted phenoxy, and halogen.

5. (original) The composition of claim 4, wherein R^1 is hydrogen and R^2 is substituted phenyl, wherein the substituents on R^2 are independently chosen from amino, (lower)alkylamino, and di(lower)alkylamino, and are located at one or more of positions 3- and 4- of the phenyl ring, in relation to the sulfonamido group.

6. (canceled)

7. (canceled)

8. (original) The composition of claim 2, wherein the compound is Pentafluorophenylsulfonamidobenzene.

9. (original) The composition of claim 3, wherein R^1 is hydrogen, and R^2 is phenyl substituted at positions 3- and 4-, in relation to the sulfonamido group, with a divalent moiety that forms a 5- or 6-membered ring together with carbons 3- and 4- of the phenyl ring.

10. (original) The composition of claim 9, wherein the divalent moiety is: $-\text{OCH}_2\text{CH}_2\text{O}-$, $-\text{OCH}_2\text{O}-$, $-\text{C}=\text{CNH}-$, or $-\text{C}=\text{NNH}-$.

11. (original) The composition of claim 10, wherein the compound is 1,2-Ethylenedioxy-4-pentafluorophenylsulfonamidobenzene,

1,2-Methylenedioxy-4-pentafluorophenylsulfonamidobenzene,
5-Pentafluorophenylsulfonamidoindazole, or
5-Pentafluorophenylsulfonamidoindole.

a' 12. (original) The composition of claim 4, wherein R^1 is hydrogen, and the substituents on R^2 are independently selected from halogen, hydroxy, lower alkyl, lower alkoxy, amino, (lower)alkylamino, and di(lower)alkylamino.

13. (original) The composition of claim 12, wherein the substituents on R^2 are independently selected from bromo, chloro, fluoro, hydroxy, methoxy, ethoxy, amino, and dimethylamino.

14. (original) The composition of claim 13, wherein the substituents on R^2 are independently selected from bromo, chloro, fluoro, hydroxy, methoxy, and ethoxy.

15. (original) The composition of claim 12, wherein the substituents on R^2 are at one or more of positions 3- and 4- of the phenyl ring, in relation to the sulfonamido group.

16. (original) The composition of claim 15, wherein R^2 is monosubstituted phenyl.

17. (canceled)

18. (original) The composition of claim 16, wherein the compound is
3-Chloro-1-pentafluorophenylsulfonamidobenzene, or
4-Chloro-1-pentafluorophenylsulfonamidobenzene.

19. (original) The composition of claim 15, wherein R^2 is disubstituted phenyl.

20. (canceled)

21. (canceled)

22. (canceled)

23. (canceled)

24. (canceled)

25. (canceled)

26. (canceled)

27. (original) The composition of claim 12, wherein R^2 is a trisubstituted phenyl.

28. (canceled)

29. (canceled)

30. (original) The composition of claim 1, wherein in the compound of formula I,
Y is SO_2 and

Z is NR^1R^2 , where R^1 is hydrogen or lower alkyl, and R^2 is an unsubstituted or
optionally substituted naphthyl group.

31. (canceled)

32. (original) The composition of claim 4, wherein R^2 is a phenyl group
substituted by phenoxy or optionally substituted phenoxy.

33. (original) The composition of claim 32, wherein the compound is 3-Phenoxy-
1-pentafluorophenylsulfonamidobenzene.

34. (original) The composition of claim 3, wherein R^2 is a phenyl ring substituted
by a heterocyclic group at the 4- position, in relation to the sulfonamido group.

35. (canceled)

36. (original) The composition of claim 2, wherein R^1 and R^2 are covalently
joined in a moiety that forms a 5- or 6-membered heterocyclic ring with the nitrogen atom of
 NR^1R^2 .

a' 37. (original) The composition of claim 36, wherein R¹ is a -CH=CH- group linked to the 2-position of the R² phenyl group, in relation to the sulfonamido group, forming an optionally substituted indole.

38. (canceled)

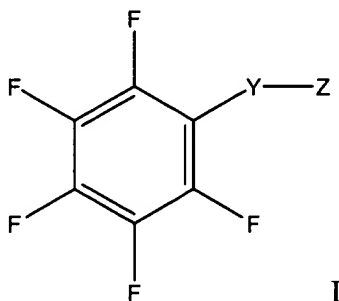
39. (original) The composition of claim 36, wherein R¹ is a -(CH₂)₃- group linked to the 2- position of the R² phenyl group, in relation to the sulfonamido group, forming an optionally substituted 1,2,3,4-tetrahydroquinoline.

40. (original) The composition of claim 39, wherein the compound is 1-pentafluorophenylsulfonyl-1,2,3,4-tetrahydroquinoline.

41. (original) The composition of claim 2, wherein R¹ is an optionally substituted (C2-C10)alkyl or optionally substituted (C2-C6)heteroalkyl.

42. (canceled)

43. (original) A method of treating or preventing a disease state characterized by abnormally high levels of low density lipoprotein particles or cholesterol in the blood, which method comprises administering to a mammalian subject in need thereof a therapeutically effective amount of a composition containing a compound of formula I



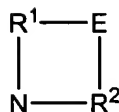
or a pharmaceutically acceptable salt thereof, wherein:

Y is -S(O)- or -S(O)₂-;

Z is -NR¹R² or -OR³; where R¹ and R² are independently selected from

a'
hydrogen,
substituted or unsubstituted (C1-C10)alkyl,
substituted or unsubstituted (C1-C10)alkoxy,
substituted or unsubstituted (C3-C6)alkenyl,
substituted or unsubstituted (C2-C6)heteroalkyl,
substituted or unsubstituted (C3-C6)heteroalkenyl,
substituted or unsubstituted (C3-C6)alkynyl,
substituted or unsubstituted (C3-C8)cycloalkyl,
substituted or unsubstituted (C5-C7)cycloalkenyl,
substituted or unsubstituted (C5-C7)cycloalkadienyl,
substituted or unsubstituted aryl,
substituted or unsubstituted aryloxy,
substituted or unsubstituted aryl-(C3-C8)cycloalkyl,
substituted or unsubstituted aryl-(C5-C7)cycloalkenyl,
substituted or unsubstituted aryloxy-(C3-C8)cycloalkyl,
substituted or unsubstituted aryl-(C1-C4)alkyl,
substituted or unsubstituted aryl-(C1-C4)alkoxy,
substituted or unsubstituted aryl-(C1-C4)heteroalkyl,
substituted or unsubstituted aryl-(C3-C6)alkenyl,
substituted or unsubstituted aryloxy-(C1-C4)alkyl,
substituted or unsubstituted aryloxy-(C2-C4)heteroalkyl,
substituted or unsubstituted heteroaryl,
substituted or unsubstituted heteroaryloxy,
substituted or unsubstituted heteroaryl-(C1-C4)alkyl,
substituted or unsubstituted heteroaryl-(C1-C4)alkoxy,
substituted or unsubstituted heteroaryl-(C1-C4)heteroalkyl,
substituted or unsubstituted heteroaryl-(C3-C6)alkenyl,
substituted or unsubstituted heteroaryloxy-(C1-C4)alkyl, and
substituted or unsubstituted heteroaryloxy-(C2-C4)heteroalkyl,

wherein R¹ and R² may be connected by a linking group E to give a substituent of the formula



a' wherein E represents a bond, (C1-C4) alkylene, or (C1-C4) heteroalkylene, and the ring formed by R¹, E, R² and the nitrogen contains no more than 8 atoms; and where R³ is optionally substituted aryl or optionally substituted heteroaryl.

44. (original) The method of claim 43 wherein, in the compound of formula I, Y is SO₂ and Z is NR¹R²; where R² is optionally substituted aryl or optionally substituted heteroaryl.

45. (original) The method of claim 44, wherein R¹ is hydrogen or lower alkyl, R² is optionally substituted phenyl, and there is no linking group E between R¹ and R².

46. (original) The method of claim 45, wherein R¹ is hydrogen or methyl, and the substituents on R² are independently chosen from lower alkyl, hydroxy, lower alkoxy, amino, amino optionally substituted with one or two lower alkyls, optionally substituted arylamino, optionally substituted heteroarylamino, optionally substituted phenoxy, and halogen.

47. (canceled)

48. (original) The method of claim 47, wherein the compound is chosen from:
4-(N,N-Dimethylamino)-1-pentafluorophenylsulfonamidobenzene,
3-(N,N-Dimethylamino)-1-pentafluorophenylsulfonamidobenzene,
1,2-Ethylenedioxy-4-pentafluorophenylsulfonamidobenzene,
2-Hydroxy-1-methoxy-4-pentafluorophenylsulfonamidobenzene,
2-Fluoro-1-methoxy-4-pentafluorophenylsulfonamidobenzene,
4-Methoxy-1-pentafluorophenylsulfonamidobenzene,
3-Hydroxy-1-pentafluorophenylsulfonamidobenzene,
4-Hydroxy-1-pentafluorophenylsulfonamidobenzene,
1,2-Dimethyl-4-pentafluorophenylsulfonamidobenzene,
5-Pentafluorophenylsulfonamidoindole,

a'

4-(N,N-Dimethylamino)-1-(N-methylpentafluorophenylsulfonamido)benzene,
4-Ethoxy-1-pentafluorophenylsulfonamidobenzene,
3-Methoxy-1-pentafluorophenylsulfonamidobenzene,
2-Bromo-1-methoxy-4-pentafluorophenylsulfonamidobenzene,
2-Chloro-1-methoxy-4-pentafluorophenylsulfonamidobenzene,
2-Bromo-3-hydroxy-4-methoxy-1-pentafluorophenylsulfonamidobenzene,
2-Bromo-4-methoxy-5-hydroxy-1-pentafluorophenylsulfonamidobenzene,
1-Bromo-4-fluoro-5-methoxy-2-pentafluorophenylsulfonamidobenzene,
2-Hydroxy-1-methoxy-4-pentafluorophenylsulfonamidobenzene; monosodium salt,
2-Hydroxy-1-methoxy-4-pentafluorophenylsulfonamidobenzene; monopotassium salt,
2-Fluoro-1-methoxy-4-pentafluorophenylsulfonamidobenzene; sodium salt,
2-Fluoro-1-methoxy-4-pentafluorophenylsulfonamidobenzene; potassium salt,
4-Chloro-1-pentafluorophenylsulfonamidobenzene, and
3-Amino-4-methoxy-1-pentafluorophenylsulfonamidobenzene.

49. (original) The method of claim 48, wherein the compound is:

2-Hydroxy-1-methoxy-4-pentafluorophenylsulfonamidobenzene,
2-Fluoro-1-methoxy-4-pentafluorophenylsulfonamidobenzene,
4-Methoxy-1-pentafluorophenylsulfonamidobenzene,
1-,2-Dimethyl-4-pentafluorophenylsulfonamidobenzene,
5-Pentafluorophenylsulfonamidoindole,
4-Ethoxy-1-pentafluorophenylsulfonamidobenzene,
2-Bromo-1-methoxy-4-pentafluorophenylsulfonamidobenzene,
2-Chloro-1-methoxy-4-pentafluorophenylsulfonamidobenzene,
2-Bromo-3-hydroxy-4-methoxy-1-pentafluorophenylsulfonamidobenzene,
2-Bromo-4-methoxy-5-hydroxy-1-pentafluorophenylsulfonamidobenzene,
1-Bromo-4-fluoro-5-methoxy-2-pentafluorophenylsulfonamidobenzene,
2-Hydroxy-1-methoxy-4-pentafluorophenylsulfonamidobenzene; monosodium salt,
2-Hydroxy-1-methoxy-4-pentafluorophenylsulfonamidobenzene; monopotassium salt,
2-Fluoro-1-methoxy-4-pentafluorophenylsulfonamidobenzene; sodium salt,
2-Fluoro-1-methoxy-4-pentafluorophenylsulfonamidobenzene; potassium salt, or

3-Amino-4-methoxy-1-pentafluorophenylsulfonamidobenzene.

50. (canceled)

51. (canceled)

52. (canceled)

53. (canceled)

54. (original) The method of claim 43, wherein the disease state is atherosclerosis.

55. (original) The method of claim 43, wherein the disease state is pancreatitis.

56. (original) The method of claim 43, wherein the disease state is hypercholesterolemia.

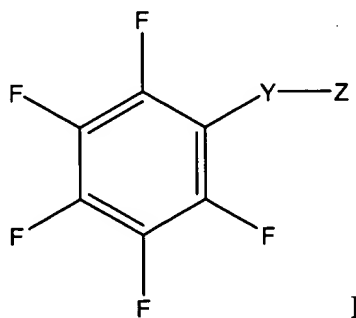
57. (original) The method of claim 43, wherein the disease state is hyperlipoproteinemia.

58. (original) The method of claim 43, wherein the composition is administered orally.

59. (original) The method of claim 43, wherein the subject is human.

60. (original) The method of claim 43, wherein the composition is administered in combination with a therapeutically effective amount of a hypolipemic agent or a hypocholesterolemic agent that is not represented by formula I.

61. (amended) A compound having the formula I:



or a pharmaceutically acceptable salt thereof, wherein:

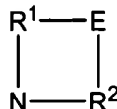
Y is -S(O)- or -S(O₂)-; and

Z is NR¹R², wherein R² is an optionally substituted aryl or heteroaryl group, and R¹ is

*a*¹ selected from

hydrogen,
substituted or unsubstituted (C1-C10)alkyl,
substituted or unsubstituted (C1-C10)alkoxy,
substituted or unsubstituted (C3-C6)alkenyl,
substituted or unsubstituted (C2-C6)heteroalkyl,
substituted or unsubstituted (C3-C6)heteroalkenyl,
substituted or unsubstituted (C3-C6)alkynyl,
substituted or unsubstituted (C3-C8)cycloalkyl,
substituted or unsubstituted (C5-C7)cycloalkenyl,
substituted or unsubstituted (C5-C7)cycloalkadienyl,
substituted or unsubstituted aryl,
substituted or unsubstituted aryloxy,
substituted or unsubstituted aryl-(C3-C8)cycloalkyl,
substituted or unsubstituted aryl-(C5-C7)cycloalkenyl,
substituted or unsubstituted aryloxy-(C3-C8)cycloalkyl,
substituted or unsubstituted aryl-(C1-C4)alkyl,
substituted or unsubstituted aryl-(C1-C4)alkoxy,
~~substituted or unsubstituted aryl-(C1-C4)heteroalkyl,~~
substituted or unsubstituted aryl-(C3-C6)alkenyl,
substituted or unsubstituted aryloxy-(C1-C4)alkyl,
substituted or unsubstituted aryloxy-(C2-C4)heteroalkyl,
substituted or unsubstituted heteroaryl,
substituted or unsubstituted heteroaryloxy,
substituted or unsubstituted heteroaryl-(C1-C4)alkyl,
substituted or unsubstituted heteroaryl-(C1-C4)alkoxy,
substituted or unsubstituted heteroaryl-(C1-C4)heteroalkyl,

a' substituted or unsubstituted heteroaryl-(C3-C6)alkenyl,
substituted or unsubstituted heteroaryloxy-(C1-C4)alkyl, and
substituted or unsubstituted heteroaryloxy-(C2-C4)heteroalkyl,
wherein R¹ and R² may be connected by a linking group E to give a substituent of the formula



wherein E represents a bond, (C1-C4) alkylene, or (C1-C4) heteroalkylene, and the ring formed by R¹, E, R² and the nitrogen contains no more than 8 atoms; provided that: in the case that Y is -S(O₂)-, and R¹ is hydrogen or methyl, then R² is substituted phenyl or heteroaryl group;

in the case that Y is -S(O₂)-, and R² is a ring system chosen from 1-naphthyl, 5-quinolyl, or 4-pyridyl, then either R¹ is not hydrogen or R² is substituted by at least one substituent that is not hydrogen;

in the case that Y is -S(O₂)-, R² is phenyl, and R¹ is a propylene unit attaching the nitrogen of -NR¹R²- to the 2- position of the phenyl ring in relation to the sulfonamido group to form a 1-,2-,3-,4-tetrahydroquinoline system, and one or more of the remaining valences on the bicyclic system so formed is substituted with at least one substituent that is not hydrogen;

in the case that Y is -S(O₂)- and R² is phenyl substituted with 3-(1-hydroxyethyl), 3-dimethylamino, 4-dimethylamino, 4-phenyl, 3-hydroxy, 3-hydroxy-4-diethylaminomethyl, 3,4-methylenedioxy, 3,4-ethylenedioxy, 2-(1-pyrrolyl), or 2-methoxy 4-(1-morpholino), then either R¹ is not hydrogen or when R¹ is hydrogen, one or more of the remaining valences on the phenyl ring of R² is substituted with a substituent that is not hydrogen;

in the case that Y is -S(O₂)- and R² is 2-methylbenzothiazol-5-yl, 6-hydroxy-4-methylpyrimidin-2-yl, 3-carbomethoxypyrazin-2-yl, 5-carbomethoxypyrazin-2-yl, 4-carboethoxy-1-phenylpyrazol-5-yl, 3-methylpyrazol-5-yl, 4-chloro-2-methylthiopyrimidin-6-yl, 2-trifluoromethyl-1-,3-,4-thiadiazol-5-yl, 5,6,7,8-tetrahydro-2-naphthyl, 4-methylthiazol-2-yl, 6,7-dihydroindan-5-yl, 7-chloro-5-methyl-1,8-naphthyridin-2-yl, 5,7-dimethyl-1,8-naphthyridin 2-yl, or 3-cyanopyrazol-4-yl, then R¹ is a group other than hydrogen;
wherein said compound has pharmacological activity.

a' 62. (original) The compound of claim 61, wherein R^1 is hydrogen or lower alkyl, Y is $-S(O_2)-$, and

there is no linking group E between R^1 and R^2 .

63. (original) The compound of claim 62, wherein R^1 is hydrogen or methyl and R^2 is substituted phenyl, wherein the substituents on R^2 , ranging in number from one to four, are independently chosen from lower alkyl, hydroxy, lower alkoxy, amino optionally substituted with one or two lower alkyls, optionally substituted arylamino, optionally substituted heteroarylamino, optionally substituted phenoxy, and halogen.

64. (original) The compound of claim 63, wherein R^1 is hydrogen and R^2 is substituted phenyl, wherein the substituents on R^2 are independently chosen from amino, (lower)alkylamino, and di(lower)alkylamino, and are located at one or more of positions 3- and 4- of the phenyl ring, in relation to the sulfonamido group.

65. (original) The compound of claim 64, wherein the compound is 4-(*N,N*-Diethylamino)-1-pentafluorophenylsulfonamidobenzene, or 4-Amino-1-pentafluorophenylsulfonamidobenzene.

66. (original) The compound of claim 62, wherein R^1 is hydrogen, and R^2 is phenyl substituted at positions 3- and 4-, in relation to the sulfonamido group, with a divalent moiety that forms a 5- or 6- membered ring together with carbons 3- and 4- of the phenyl ring.

67. (original) The compound of claim 66, wherein the divalent moiety is: - $C=CNH-$, or $-C=NNH-$.

68. (canceled)

69. (original) The compound of claim 63, wherein R^1 is hydrogen, and the substituents on R^2 are independently selected from halogen, hydroxy, lower alkyl, lower alkoxy, amino, (lower)alkylamino, and di(lower)alkylamino.

a! 70. (original) The compound of claim 69, wherein the substituents on R² are independently selected from bromo, chloro, fluoro, hydroxy, methoxy, ethoxy, amino, or dimethylamino:

71. (original) The compound of claim 70, wherein the substituents on R² are independently selected from bromo, chloro, fluoro, hydroxy, methoxy, and ethoxy.

72. (original) The compound of claim 71, wherein the substituents on R² are at one or more of positions 3- and 4- of the phenyl ring, in relation to the sulfonamido group.

73. (original) The compound of claim 72, wherein R² is monosubstituted phenyl.

74. (canceled)

75. (canceled)

76. (original) The compound of claim 72, wherein R² is disubstituted phenyl.

77. (canceled).

78. (canceled)

79. (original) The compound of claim 77, wherein the compound is
2-Hydroxy-1-methoxy-4-pentafluorophenylsulfonamidobenzene, or
2-Hydroxy-1-methoxy-4-pentafluorophenylsulfonamidobenzene, monosodium salt.

80. (original) The compound of claim 78, wherein the compound is
2-Fluoro 1-methoxy-4-pentafluorophenylsulfonamidobenzene, or
2-Fluoro 1-methoxy-4-pentafluorophenylsulfonamidobenzene, sodium salt.

81. (canceled)

82. (canceled)

83. (original) The compound of claim 72, wherein R² is trisubstituted phenyl.

a' 84. (original) The compound of claim 78, wherein the compound is 2-Bromo-1-methoxy-4-pentafluorophenylsulfonamidobenzene, or 2-Chloro-1-methoxy-4-pentafluorophenylsulfonamidobenzene.

85. (canceled)

86. (canceled)

87. (original) The compound of claim 62, wherein R^2 is a phenyl ring substituted by a heterocyclic group at the 4- position, in relation to the sulfonamido group.

88. (canceled)

89. (original) The compound of claim 61, wherein R^1 and R^2 are covalently joined in a moiety that forms a 5- or 6-membered heterocyclic ring with the nitrogen atom of NR^1R^2 .

90. (original) The compound of claim 89, wherein R^1 is a $-CH=CH-$ group linked to the 2- position of the R^2 phenyl group, in relation to the sulfonamido group, forming an optionally substituted indole.

91. (canceled)

92. (original) The compound of claim 89, wherein R^1 is a $-(CH_2)_3-$ group linked to the 2- position of the R^2 phenyl group, in relation to the sulfonamido group, forming an optionally substituted 1,2,3,4-tetrahydroquinoline.

93. (canceled)

94. (canceled)